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                 INPADOC replaced by INPADOCDB on STN
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        APR 30
                New CAS web site launched
NEWS 12
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        MAY 08
NEWS 13
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NEWS 14
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                 fields
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                 TOXCENTER enhanced with BIOSIS reload
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                CA/CAplus enhanced with IPC reclassification in Japanese
NEWS 18
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                 patents
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         JUN 27
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                 STN Express, Version 8.2, now available
         JUN 29
NEWS 21
NEWS 22 JUL 02 LEMBASE coverage updated
NEWS 23 JUL 02 LMEDLINE coverage updated
                 SCISEARCH enhanced with complete author names
NEWS 24
         JUL 02
         JUL 02 CHEMCATS accession numbers revised
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                 CA/CAplus enhanced with utility model patents from China
         JUL 02
NEWS 26
                 CAplus enhanced with French and German abstracts
NEWS 27
         JUL 16
NEWS 28 JUL 18 CA/CAplus patent coverage enhanced
NEWS EXPRESS
              29 JUNE 2007: CURRENT WINDOWS VERSION IS V8.2,
              CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 05 JULY 2007.
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              For general information regarding STN implementation of IPC 8
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FILE 'BIOSIS' ENTERED AT 13:29:13 ON 23 JUL 2007 Copyright (c) 2007 The Thomson Corporation

=> retinoid

L1 30662 RETINOID

=> HCV

L2 38087 HCV

=> 11 and L2

L3 19 L1 AND L2

=> selnium

1 SELNIUM L4

=> selenium

L5 115006 SELENIUM

=> L5 and L2

27 L5 AND L2 L6

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L7 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN.

ACCESSION NUMBER:

2005:1331259 CAPLUS 144:64327

DOCUMENT NUMBER: TITLE:

Use of selenium or a selenium salt and a retinoid acid or a retinoid in the treatment of viral hepatitis C

Herget, Thomas; Klebl, Bert INVENTOR(S): GPC Biotech A.-G., Germany

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND DATE	APPLICATION NO.	DATE			
WO 2005120479	A1 20051222	WO 2005-EP6226	20050609			
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GE, GH, G	i, HR, HU, ID, IL,	IN, IS, JP, KE, KG, KM,	KP, KR, KZ,			
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PRIORITY APPLN. INFO.:
                                                       US 2004-578161P
                                                                                P 20040609 '
      The present invention relates to combination therapies comprising at least
AB
      one retinoid or retinoid agonist together with
      selenium or a selenium salt particularly useful in
      conjunction with conventional antiviral therapeutics which are
      synergistically effective against Hepatitis C virus (HCV)
      infections. In particular, the present invention relates to the synergism
      between compds. capable of activating or upregulating the gastrointestinal
      form of glutathione peroxidase for prophylaxis and/or treatment of
      HCV infections, administered in combination therapies with
      interferons. The combinations disclosed have proven surprisingly
      effective even in patients unresponsive to interferon/ribavirin therapies.
                                      THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
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                                      RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
      ANSWER 2 OF 4
                       CAPLUS COPYRIGHT 2007 ACS on STN
                               2004:633154 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                               141:167729
TITLE:
                               Gastrointestinal glutathione peroxidase as therapeutic
                               target for treatment of HCV infection,
                               methods of treating HCV infection, and
                               compounds useful therefor
                               Herget, Thomas; Cotten, Matthew; Obert, Sabine; Klebl,
INVENTOR(S):
                               Bert
PATENT ASSIGNEE(S):
                               Germany
                               U.S. Pat. Appl. Publ., 24 pp., Cont.-in-part of U.S.
SOURCE:
                               Pat. Appl. 2003 180,719.
                               CODEN: USXXCO
DOCUMENT TYPE:
                               Patent
LANGUAGE:
                               English
FAMILY ACC. NUM. COUNT:
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                                                       US 2003-723719
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

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                                                       US 2001-283345P
PRIORITY APPLN. INFO.:
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                                                                                A 20021129
                                                                               P 20021203
                                                       US 2002-430367P
                                                       US 2003-342054
                                                                                A2 20030114
      The present invention relates to the human cellular protein glutathione
AB
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peroxidase-gastrointestinal as a target for medical intervention against

Hepatitis C virus (HCV) infections. Furthermore, the present invention relates to a method for the detection of compds. useful for prophylaxis and/or treatment of hepatitis C virus infections and a method for detecting hepatitis C virus infections in an individual or in cells. Also compns., compds., nucleic acid mols. (such as aptamers), mono- or polyclonal antibodies are disclosed which are effective for the treatment of HCV infections, and methods for prophylaxis and/or treatment of hepatitis C virus infections or for the regulation of hepatitis C virus production are disclosed. The inventors designed a randomized, single-blinded clin. study to test the safety, tolerability, and efficacy of all-trans retinoic acid alone or in combination with pegylated  $\alpha$  interferon in patients with chronic hepatitis C. The therapy regimens include: Vesanoid (orally administered all-trans retinoic acid compound, Hoffman-La Roche); Pegasys (slow-release pegylated interferon α2a, Hoffman-La Roche); and selen 30 ALLACT (supplement containing selenium and ALLACT composed of garlic powder and Lactobacillus bulgaricus).

L7 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:490732 CAPLUS

DOCUMENT NUMBER:

141:42933

TITLE:

Formulations useful against hepatitis C virus

infections

INVENTOR(S):

Herget, Thomas; Klebl, Bert

PATENT ASSIGNEE(S):

Axxima Pharmaceuticals A.-G., Germany

SOURCE:

PCT Int. Appl., 72 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PRIORITY APPLN. INFO.:
                                           DE 2002-10255861
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                                           US 2002-430367P
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                                            US 2003-446246P
                                                                  20030211
                                           WO 2003-EP13514
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AB The present invention relates generally to chemical compds. and substances which are effective against Hepatitis C virus (HCV) infections.

Moreover, the present invention relates to compns. comprising said compds. and/or substances, to methods for preventing HCV infections as well use of the compds. and/or substances for the preparation of compns. useful

for the prophylaxis and/or treatment of HCV infections. Useful compds. and substances according to the invention are selenium, selenium salts, Vitamin D3 and retinoids, like all trans retinoic acid and salts thereof, C1-C10 alkyl amide of all trans retinoic acid and salts thereof, C1-C10 alkyl esters of all trans retinoic acid and salts thereof, 9-cis retinoic acid and salts thereof, C1-C10 alkyl amide of 9-cis retinoic acid and salts thereof, C1-C10 alkyl esters of 9-cis retinoic acid and salts thereof, C1-C10 alkyl esters of 9-cis retinoic acid and salts thereof, (E)-4-[2-(5,6,7,8-tetrahydro-5,5,8,8-tetra methyl-2-naphthalenyl-1)-propenyl] benzoic acid (TTNPB), (4-[5,6,7,8-tetrahydro-5,5,8,8-tetramethyl-2-naphthalenyl] carboxamido) benzoic acid (AM-580), N-(4-hydroxyphenyl) retinamide (4-HPR), and 6-[3-(1-adamantyl)-4-hydroxyphenyl]-2-naphthalene carboxylic acid (AHPN).

L7 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

PATENT ASSIGNEE(S):

2003:757185 CAPLUS

DOCUMENT NUMBER:

139:271014

TITLE:

Human cellular protein gastrointestinal glutathione

peroxidase as target for medical intervention against

hepatitis C virus infections

INVENTOR(S):

Herget, Thomas; Cotten, Matthew; Obert, Sabine

Germany

SOURCE:

U.S. Pat. Appl. Publ., 23 pp., Cont.-in-part of Appl.

No. PCT/EP02/04167.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

P	PATENT NO.						KIND DATE			;	APPL	ICAT:	DATE							
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										1	WO 2	002-	EP41	67		A2 2	0020	415		
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The present invention relates to the human cellular protein glutathione peroxidase-gastrointestinal as a target for medical intervention against Hepatitis C virus (HCV) infections. Furthermore, the present invention relates to a method for the detection of compds. useful for prophylaxis and/or treatment of Hepatitis C virus infections and a method for detecting Hepatitis C virus infections in an individual or in cells. Also compns., compds., nucleic acid mols. (such as aptamers), mono- or polyclonal antibodies are disclosed which are effective for the treatment of HCV infections, and methods for prophylaxis and/or treatment of Hepatitis C virus infections or for the regulation of Hepatitis C virus production are disclosed.

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L8
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=> ribavirin
           10661 RIBAVIRIN
L9
=> L8 and L9
L10
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=> L10 and L3
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L11
=> L10 and L6
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MISSING OPERATOR L11 IBIB
The search profile that was entered contains terms or
nested terms that are not separated by a logical operator.
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L11 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN
                               2005:1331259 CAPLUS
ACCESSION NUMBER:
                               144:64327
DOCUMENT NUMBER:
                               Use of selenium or a selenium salt and a
TITLE:
                               retinoid acid or a retinoid in the
                               treatment of viral hepatitis C
                               Herget, Thomas; Klebl, Bert
INVENTOR(S):
                               GPC Biotech A.-G., Germany
PATENT ASSIGNEE(S):
                               PCT Int. Appl., 58 pp.
SOURCE:
                               CODEN: PIXXD2
DOCUMENT TYPE:
                               Patent
                               English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
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      WO 2005120479
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                MR, NE, SN, TD, TG
                                                       US 2004-578161P
                                                                                P 20040609
PRIORITY APPLN. INFO.:
      The present invention relates to combination therapies comprising at least
      one retinoid or retinoid agonist together with
      selenium or a selenium salt particularly useful in conjunction with
       conventional antiviral therapeutics which are synergistically effective
       against Hepatitis C virus (HCV) infections. In particular, the
      present invention relates to the synergism between compds. capable of
       activating or upregulating the gastrointestinal form of glutathione
      peroxidase for prophylaxis and/or treatment of HCV infections,
       administered in combination therapies with interferons. The
       combinations disclosed have proven surprisingly effective even in patients
```

unresponsive to interferon/ribavirin therapies.

=> interferon

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:633154 CAPLUS

DOCUMENT NUMBER: 141:167729

TITLE: Gastrointestinal glutathione peroxidase as therapeutic

target for treatment of HCV infection, methods of treating HCV infection, and

compounds useful therefor

INVENTOR(S): Herget, Thomas; Cotten, Matthew; Obert, Sabine; Klebl,

Bert

PATENT ASSIGNEE(S): Germany

SOURCE: U.S. Pat. Appl. Publ., 24 pp., Cont.-in-part of U.S.

Pat. Appl. 2003 180,719.

CODEN: USXXCO

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

US 2004152073 A1 20040805 US 2003-723719 20031126 WO 2002084294 A2 20021024 WO 2002-EP4167 20020415 WO 2002084294 A3 20031030 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, TR, TR, TT, TZ, TM, TN, TR, TT, TZ, TM, TM, TN, TR, TT, TZ, TM, TM, TN, TN, TR, TT, TZ, TM, TM, TN, TN, TN, TN, TR, TT, TZ, TM, TM, TN, TN, TN, TN, TN, TN, TN, TN, TN, TN
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DE 10255861 A1 20040617 DE 2002-10255861 20021129
US 2003180719 A1 20030925 US 2003-342054 20030114
PRIORITY APPLN. INFO.: US 2001-283345P P 20010413
WO 2002-EP4167 A2 20020415
DE 2002-10255861 A 20021129
US 2002-430367P P 20021203
US 2003-342054 A2 20030114

The present invention relates to the human cellular protein glutathione AB peroxidase-gastrointestinal as a target for medical intervention against Hepatitis C virus (HCV) infections. Furthermore, the present invention relates to a method for the detection of compds. useful for prophylaxis and/or treatment of hepatitis C virus infections and a method for detecting hepatitis C virus infections in an individual or in cells. Also compns., compds., nucleic acid mols. (such as aptamers), mono- or polyclonal antibodies are disclosed which are effective for the treatment of HCV infections, and methods for prophylaxis and/or treatment of hepatitis C virus infections or for the regulation of hepatitis C virus production are disclosed. The inventors designed a randomized, single-blinded clin. study to test the safety, tolerability, and efficacy of all-trans retinoic acid alone or in combination with pegylated  $\boldsymbol{\alpha}$ interferon in patients with chronic hepatitis C. The therapy regimens include: Vesanoid (orally administered all-trans retinoic acid compound, Hoffman-La Roche); Pegasys (slow-release pegylated interferon  $\alpha$ 2a, Hoffman-La Roche); and selen 30 ALLACT (supplement containing selenium and ALLACT composed of garlic powder and Lactobacillus bulgaricus).

ACCESSION NUMBER:

2004:490732 CAPLUS

DOCUMENT NUMBER:

141:42933

TITLE:

Formulations useful against hepatitis C virus

infections

INVENTOR(S):

Herget, Thomas; Klebl, Bert

PATENT ASSIGNEE(S):

Axxima Pharmaceuticals A.-G., Germany

SOURCE:

PCT Int. Appl., 72 pp. CODEN: PIXXD2

Patent

DOCUMENT TYPE: LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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                                  20040617
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     WO 2004050101
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              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
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             BY, KG, KZ, MD, RU; TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                  20060427
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     US 2006151574
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PRIORITY APPLN. INFO.:
                                                                   P 20021203
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                                                                   A 20030207
                                               US 2003-446246P
                                                                    P 20030211
                                               WO 2003-EP13514
                                                                     W 20031201
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The present invention relates generally to chemical compds. and substances AΒ which are effective against Hepatitis C virus (HCV) infections. Moreover, the present invention relates to compns. comprising said compds. and/or substances, to methods for preventing HCV infections as well use of the compds. and/or substances for the preparation of compns. useful for the prophylaxis and/or treatment of HCV infections. Useful compds. and substances according to the invention are selenium, selenium salts, Vitamin D3 and retinoids, like all trans retinoic acid and salts thereof, C1-C10 alkyl amide of all trans retinoic acid and salts thereof, C1-C10 alkyl esters of all trans retinoic acid and salts thereof, 9-cis retinoic acid and salts thereof, C1-C10 alkyl amide of 9-cis retinoic acid and salts thereof, C1-C10 alkyl esters of 9-cis retinoic acid and salts thereof, (E)-4-[2-(5,6,7,8-tetrahydro-5,5,8,8-tetra methyl-2-naphthalenyl-1)-propenyl] benzoic acid (TTNPB), (4-[5,6,7,8-tetrahydro-5,5,8,8-tetramethyl-2-naphthalenyl] carboxamido) benzoic acid (AM-580), N-(4-hydroxyphenyl) retinamide (4-HPR), and 6-[3-(1-adamanty1)-4-hydroxypheny1]-2-naphthalene carboxylic acid (AHPN).

## => D L12 IBIB ABS 1-8

L12 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: • 2005:1331259 CAPLUS

DOCUMENT NUMBER:

144:64327

TITLE:

Use of selenium or a selenium salt

and a retinoid acid or a retinoid in the treatment of

viral hepatitis C

INVENTOR(S):

PATENT ASSIGNEE(S):

Herget, Thomas; Klebl, Bert GPC Biotech A.-G., Germany

PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

SOURCE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	PATENT NO.					KIND DATE				APPL	ICAT:		DATE				
	WO 2005120479					A1 20051222				· · · · ·	005-1		20050609				
WO	2005		-														
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		NG,	NI,	NO,	ΝZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,
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		EE,	ES,	FI,	FR,	GB,	GR,	ΗU,	ΙE,	IS,	IT,	LT,	LU,	MC,	ΝL,	PL,	PT,
		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,
		MR,	ΝĒ,	SN,	TD,	TG											

PRIORITY APPLN. INFO.:

US 2004-578161P P 20040609

The present invention relates to combination therapies comprising at least one retinoid or retinoid agonist together with selenium or a selenium salt particularly useful in conjunction with conventional antiviral therapeutics which are synergistically effective against Hepatitis C virus (HCV) infections. In particular, the present invention relates to the synergism between compds. capable of activating or upregulating the gastrointestinal form of glutathione peroxidase for prophylaxis and/or treatment of HCV infections, administered in combination therapies with interferons. The combinations disclosed have proven surprisingly effective even in patients unresponsive to interferon/ribavirin therapies.

REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

8

ACCESSION NUMBER:

2004:633154 CAPLUS

DOCUMENT NUMBER:

141:167729

TITLE:

Gastrointestinal glutathione peroxidase as therapeutic

target for treatment of HCV infection, methods of treating HCV infection, and

compounds useful therefor

INVENTOR (S):

Herget, Thomas; Cotten, Matthew; Obert, Sabine; Klebl,

Bert

PATENT ASSIGNEE(S):

Germany

SOURCE:

U.S. Pat. Appl. Publ., 24 pp., Cont.-in-part of U.S.

Pat. Appl. 2003 180,719.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004152073	Al	20040805	US 2003-723719	20031126

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20020415
                                            WO 2002-EP4167
    WO 2002084294
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             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
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                                            US 2001-283345P
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PRIORITY APPLN. INFO.:
                                                                A2 20020415
                                            WO 2002-EP4167
                                            DE 2002-10255861
                                                                A 20021129
                                            US 2002-430367P
                                                                P 20021203
                                                                A2 20030114
                                            US 2003-342054
     The present invention relates to the human cellular protein glutathione
ΑB
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peroxidase-gastrointestinal as a target for medical intervention against Hepatitis C virus (HCV) infections. Furthermore, the present invention relates to a method for the detection of compds. useful for prophylaxis and/or treatment of hepatitis C virus infections and a method for detecting hepatitis C virus infections in an individual or in cells. Also compns., compds., nucleic acid mols. (such as aptamers), mono- or polyclonal antibodies are disclosed which are effective for the treatment of HCV infections, and methods for prophylaxis and/or treatment of hepatitis C virus infections or for the regulation of hepatitis C virus production are disclosed. The inventors designed a randomized, single-blinded clin. study to test the safety, tolerability, and efficacy of all-trans retinoic acid alone or in combination with pegylated  $\alpha$ interferon in patients with chronic hepatitis C. The therapy regimens include: Vesanoid (orally administered all-trans retinoic acid compound, Hoffman-La Roche); Pegasys (slow-release pegylated interferon α2a, Hoffman-La Roche); and selen 30 ALLACT (supplement containing selenium and ALLACT composed of garlic powder and Lactobacillus bulgaricus).

L12 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:490732 CAPLUS

DOCUMENT NUMBER:

141:42933

TITLE:

Formulátions useful against hepatitis C virus

infections

CODEN: PIXXD2

INVENTOR(S):

Herget, Thomas; Klebl, Bert

PATENT ASSIGNEE(S):

Axxima Pharmaceuticals A.-G., Germany

SOURCE:

PCT Int. Appl., 72 pp.

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.					KIND DATE				APPLICATION NO.						DATE			
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WO 2004050101				A2 20040617				1	WO 2003-EP13514						20031201			
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							RU,											
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     EP 1567172
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             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
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     US 2006151574
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                                            DE 2002-10255861
PRIORITY APPLN. INFO.:
                                                                P 20021203,
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                                                               P 20030211
                                            US 2003-446246P
                                                                W 20031201
                                            WO 2003-EP13514
     The present invention relates generally to chemical compds. and substances
AΒ
     which are effective against Hepatitis C virus (HCV) infections.
     Moreover, the present invention relates to compns. comprising said compds.
     and/or substances, to methods for preventing HCV infections as
     well use of the compds. and/or substances for the preparation of compns. useful
     for the prophylaxis and/or treatment of HCV infections. Useful
     compds. and substances according to the invention are selenium,
     selenium salts, Vitamin D3 and retinoids, like all trans retinoic
     acid and salts thereof, C1-C10 alkyl amide of all trans retinoic acid and
     salts thereof, C1-C10 alkyl esters of all trans retinoic acid and salts
     thereof, 9-cis retinoic acid and salts thereof, C1-C10 alkyl amide of
     9-cis retinoic acid and salts thereof, C1-C10 alkyl esters of 9-cis
     retinoic acid and salts thereof, (E)-4-[2-(5,6,7,8-tetrahydro-5,5,8,8-
     tetra methyl-2-naphthalenyl-1)-propenyl] benzoic acid (TTNPB),
     (4-[5,6,7,8-tetrahydro-5,5,8,8-tetramethyl-2-naphthalenyl] carboxamido)
     benzoic acid (AM-580), N-(4-hydroxyphenyl) retinamide (4-HPR), and
     6-[3-(1-adamantyl)-4-hydroxyphenyl]-2-naphthalene carboxylic acid (AHPN).
L12 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN
                         2004:232965 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         140:368141
                         Urinary nitrite/nitrate concentrations and total
TITLE:
                         antioxidant capacity in patients with chronic
                         hepatitis C in therapy with interferon and
                         ribavirin
                         Stanzial, A. M.; Benoni, G.; Cuzzolin, L.; Gabrielli,
AUTHOR(S):
                         G. B.; Pasino, M.; Perfetti, P.; Corrocher, R.
                         Department of Clinical & Experimental Medicine,
CORPORATE SOURCE:
                         University of Verona, Italy
                        . Journal of Chemotherapy (Firenze, Italy) (2003),
SOURCE:
                         15(6), 584-590
                         CODEN: JCHEEU; ISSN: 1120-009X
                         E.I.F.T. srl
PUBLISHER:
DOCUMENT TYPE:
                         Journal
                         English
LANGUAGE:
     In this study we tried to elucidate the role of nitric oxide (NO) in
AB
     chronic hepatitis C in relation to antioxidant status, since the
     mechanisms by which hepatitis C virus (HCV) causes cell damage
     and the factors underlying its resistance to therapy are not well
     understood. Before and after one and six months of therapy with \alpha-
     interferon and ribavirin, we measured nitrite/nitrate
     urinary levels, total antioxidant capacity and selenium serum
     concns. in 14 patients with chronic hepatitis C and in 9 healthy subjects.
     Before therapy, mean urinary nitrite/nitrate levels of patients were not
     different from those of healthy subjects, but after a 6-mo treatment with
     \alpha- interferon and ribavirin, these NO metabolites
     were higher in virol. neg. patients (responders). Moreover, while no
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BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,

changes in selenium were observed in all patients, total antioxidant capacity was significantly higher in non-responders and well correlated with hyperuricemia (due to cell damage) observed in these subjects. Instead, uric acid decreased as free mol. in serum in responders, while we found the excretion of high NO levels as nitrite/nitrate. Our data allow us to hypothesize a role for NO as predictive of the success of therapy, since nitrite/nitrate increase in the urine of some patients precedes disappearance of the virus observed at the end of therapy.

REFERENCE COUNT:

THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

37

ACCESSION NUMBER:

2003:222329 CAPLUS

DOCUMENT NUMBER:

138:231706

TITLE:

HCV combination therapy with

ribavirin and antioxidant

INVENTOR(S):

Brass, Clifford A.

PATENT ASSIGNEE(S): SOURCE:

Schering Corporation, USA U.S. Pat. Appl. Publ., 14 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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	US	2003	0550	13		A1						002-2						
	WO	2003	0244	61		A1		2003	0327	Ţ	WO 2	002-T	JS29	576		20	00209	918
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			FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	SK,	TR,	BF,	ВJ,	CF,
								GQ,										
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										1	WO 2	002-1	US29	576	7	1 2	0020	918
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Methods of treating patients having susceptible viral infections, especially ABchronic hepatitis C infection by administering to said patient a therapeutically effective amount of a combination therapy of interferon-alfa and ribavirin for a time sufficient to lower HCV-RNA in association with a therapeutically effective amount of an antioxidant therapy comprising S-adenosyl methionine, preferably S-adenosyl L-methionine, for a time sufficient to ameliorate ribavirin-related hemolysis are disclosed.

L12 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:755216 CAPLUS

DOCUMENT NUMBER:

133:317537

TITLE:

Hepatitis C virus (HCV) combination therapy,

containing ribavirin in association with

antioxidants

INVENTOR(S):

Brass, Clifford A.; Glue, Paul W.; Piken, Edward

PATENT ASSIGNEE(S): SOURCE:

Schering Corporation, USA Eur. Pat. Appl., 16 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

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APPLICATION NO.
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                                20001025 EP 2000-303246
    EP 1046399
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                                            CA 2000-2306039
                                          WO 2000-US10240
    WO 2000062799
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                                20001026
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             IS, JP, KG, KR, KZ, LC, LK, LR, LT, LU, LV, MA, MD, MG, MK, MN,
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                                         BR 2000-9840
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                                            US 1999-294687
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PRIORITY APPLN. INFO.:
                                                               W 20000418
                                            WO 2000-US10240
     Methods are disclosed for treating patients having susceptible viral
AB
     infections, especially chronic hepatitis C infection, by administering to the
     patient a therapeutically effective amount of a combination therapy of
     interferon-\alpha and ribavirin for a time sufficient
     to lower HCV-RNA in association with a therapeutically effective
     amount of an antioxidant for a time sufficient to ameliorate
     ribavirin-related hemolysis.
                               THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
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                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L12 ANSWER 7 OF 8 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
                    2006:210736 BIOSIS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                    PREV200600212465
                    All-trans-retinoic acid for treatment of patients with
TITLE: .
                    chronic hepatitis C and non-response to interferon
                    alfa/ribavirin.
                    Becher, Wulf O.; Wallasch, Christian; Herget, T.; Klebl, B.
AUTHOR(S):
                    M.; Galle, Peter R.; Strand, D.
                    Gastroenterology, (APR 2005) Vol. 128, No. 4, Suppl. 2, pp.
SOURCE:
                    A697-A698.
                    Meeting Info.: Annual Meeting of the American-
                    Gastroenterological-Association/Digestive-Disease-Week.
                    Chicago, IL, USA. May 14 -19, 2005. Amer Gastroenterol
                    Assoc.
                    CODEN: GASTAB. ISSN: 0016-5085.
DOCUMENT TYPE:
                 Conference; (Meeting)
                    Conference; Abstract; (Meeting Abstract)
                    English
LANGUAGE:
                    Entered STN: 29 Mar 2006
ENTRY DATE:
                    Last Updated on STN: 29 Mar 2006
     Introduction: In vitro studies, submitted in parallel by Herget et al,
AB
     have shown that all-trans retinoic acid (ATRA) induces upregulation of
     selenium dependent gastrointestinal-glutathione peroxidase in
     HCV-subgenomic RNA replicon cells leading to drastic
     downregulation of the replicon, that was further enhanced by
     interferon alfa. Based on these findings, a clinical pilot trial
     was performed in HCV non-responder patients. Methods: 20
     patients with chronic HCV infection and non-response to IFN alfa
     and ribavirin (pos. PCR at week 12) were randomly assigned to
     treatment with daily 45 mg/m2 ATRA p.o. and 30 mcg/d selenite (arm A) or
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45 mg/m2 ATRA and selenite combined with 180 mcg/week peg-interferon alfa2a (arm B). All patients had serotype-1, elevated ALT levels and 9 patients had F3 fibrosis or cirrhosis. Mean IFNa pretreatment duration was 14 months, 9 patients were Peg-IFN nonresponders. ATRA treatment was continued for 12 weeks and followed for additional 12 weeks after end of treatment (ETR). HCV RNA was assessed by quantitative real time PCR.

L12 ANSWER 8 OF 8 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

ACCESSION NUMBER:

2004:167903 BIOSIS ,

DOCUMENT NUMBER:

PREV200400170221

TITLE:

Urinary nitrite/nitrate concentrations and total

antioxidant capacity in patients with chronic hepatitis C

in therapy with interferon and ribavirin

AUTHOR(S):

Stanzial, A. M.; Benoni, G. [Reprint Author]; Cuzzolin, L.;

Gabrielli, G. B.; Pasino, M.; Perfetti, P.; Corrocher, R.

CORPORATE SOURCE:

Department of Medicine and Public Health-Section of

Pharmacology, University of Verona, Policlinico G.B. Rossi,

37134, Verona, Italy

quiseppina.benoni@univr.it

SOURCE:

Journal of Chemotherapy, (December 2003) Vol. 15, No. 6,

pp. 584-590. print.

ISSN: 1120-009X (ISSN print).

DOCUMENT TYPE:

Article English

LANGUAGE: ENTRY DATE:

Entered STN: 24 Mar 2004

Last Updated on STN: 24 Mar 2004

In this study we tried to elucidate the role of nitric oxide (NO) in AΒ chronic hepatitis C in relation to antioxidant status, since the mechanisms by which hepatitis C virus (HCV) causes cell damage and the factors underlying its resistance to therapy are not well understood. Before and after one and six months of therapy with alphainterferon and ribavirin, we measured nitrite/nitrate urinary levels, total antioxidant capacity and selenium serum concentrations in 14 patients with chronic hepatitis C and in 9 healthy subjects. Before therapy, mean urinary nitrite/nitrate levels of patients were not different from those of healthy subjects, but after a 6-month treatment with alpha-interferon and ribavirin, these NO metabolites were higher in virologically negative patients (responders). Moreover, while no changes in selenium were observed in all patients, total antioxidant capacity was significantly higher in non-responders and well correlated with hyperuricemia (due to cell damage) observed in these subjects. Instead, uric acid decreased as free molecule in serum in responders, while we found the excretion of high NO levels as nitrite/nitrate. Our data allow us to hypothesize a role for NO as predictive of the success of therapy, since nitrite/nitrate increase in the urine of some patients precedes disappearance of the virus observed at the end of therapy.

L6 ANSWER 24 OF 27 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 1999:478130 BIOSIS DOCUMENT NUMBER: PREV199900478130

TITLE: Interferon/antioxidant combination therapy for chronic

hepatitis C-A controlled pilot trial.

AUTHOR(S): Look, Markus P. [Reprint author]; Gerard, Alexandra; Rao,

Govind S.; Sudhop, Thomas; Fischer, Hans-Peter; Sauerbruch,

Tilman; Spengler, Ulrich

CORPORATE SOURCE: Department of General Internal Medicine, University of

Bonn, Sigmund-Freud-Strasse 25, 53105, Bonn, Germany

SOURCE: Antiviral Research, (Sept., 1999) Vol. 43, No. 2, pp. 113-122. print.

CODEN: ARSRDR. ISSN: 0166-3542.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 9 Nov 1999

Last Updated on STN: 9 Nov 1999

The effects of two forms of antioxidative co-therapy were analyzed in 24 AB interferon-alpha (IFN)-naive patients with chronic hepatitis C who were randomized to either receive IFN monotherapy (3 X 4.5 million units IFN-alpha 2a per week), (group A), or IFN and N-acetylcysteine (N-acetylcysteine (NAC) 1,800 mg/day) plus sodium selenite (400 mug/day) supplementation (group B), or treatment as in group B plus vitamin E (544 IU/day) (group C), over 24 weeks. Changes in histology, normalization of ALT, reduction of viral RNA, serum levels of glutathione, selenium , vitamin E, erythrocyte glutathione peroxidase, trolox equivalent antioxidative capacity (TEAC), thiobarbituric acid reactive substances (TBARS) and protein carbonyl groups were measured. Low baseline TEAC and elevated TBARS indicated increased oxidative stress before therapy, which was not affected by antioxidant supplementation. At the end of treatment complete responses were found in 3/8, 2/8 and 6/8 patients in groups A, B and C, respectively, but liver histology had not significantly improved. Vitamin E treated patients had a 2.4 greater chance (95% CI: 1.05-5.5) of obtaining a complete response and had significantly greater reduction in viral load (P = 0.028) than patients without vitamin E. Relapses, i.e. re-appearance of detectable hepatitis C virus (HCV) RNA and/or re-elevation of ALT-activity occurred in 7 out of the 11 responders within 6 months after termination of therapy (group A: 2/3, group B: 1/2 and group C: 4/6). Thus, no overall beneficial effect of antioxidant/IFN therapy was detected. However, the apparent trent towards a more favorable outcome with vitamin E supplementation warrants to further study this substance as an adjuvant to IFN therapy in chronic hepatitis C.

L6 ANSWER 25 OF 27 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

ACCESSION NUMBER: 2006:210736 BIOSIS DOCUMENT NUMBER: PREV200600212465

TITLE: All-trans-retinoic acid for treatment of patients with

chronic hepatitis C and non-response to interferon

alfa/ribavirin.

AUTHOR(S): Becher, Wulf O.; Wallasch, Christian; Herget, T.; Klebl, B.

M.; Galle, Peter R.; Strand, D.

SOURCE: Gastroenterology, (APR 2005) Vol. 128, No. 4, Suppl. 2, pp.

A697-A698.

Meeting Info.: Annual Meeting of the American-

Gastroenterological-Association/Digestive-Disease-Week. Chicago, IL, USA. May 14 -19, 2005. Amer Gastroenterol

Assoc.

CODEN: GASTAB. ISSN: 0016-5085.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 29 Mar 2006

Last Updated on STN: 29 Mar 2006

AB Introduction: In vitro studies, submitted in parallel by Herget et al, have shown that all-trans retinoic acid (ATRA) induces upregulation of selenium dependent gastrointestinal-glutathione peroxidase in HCV-subgenomic RNA replicon cells leading to drastic downregulation of the replicon, that was further enhanced by interferon alfa. Based on these findings, a clinical pilot trial was performed in HCV non-responder patients. Methods: 20 patients with chronic HCV infection and non-response to IFN alfa and ribavirin (pos. PCR at week 12) were randomly assigned to treatment with daily 45 mg/m2 ATRA p.o. and 30 mcg/d selenite (arm A) or 45 mg/m2 ATRA and selenite combined with 180 mcg/week peg-interferon alfa2a (arm B). All patients had serotype-1, elevated ALT levels and 9 patients had F3 fibrosis or cirrhosis. Mean IFNa pretreatment duration was 14 months, 9 patients were Peg-IFN nonresponders. ATRA treatment was continued for 12 weeks and followed for additional 12 weeks after end of treatment (ETR).

HCV RNA was assessed by quantitative real time PCR.

CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 15 OF 27

ACCESSION NUMBER:

1999:819471 CAPLUS

DOCUMENT NUMBER:

132:47240

TITLE:

Process for the in vitro replication of HCV

INVENTOR(S):

Rumin, Sylvie; Inchauspe, Genevieve; Trepo, Christian;

Gripon, Philippe

PATENT ASSIGNEE(S):

Institut National De La Sante Et De La Recherche

Medicale I.N.S.E.R.M., Fr.

SOURCE:

PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 9967362	A1 19991229	WO 1999-EP4337	19990623
W: CA, JP, US			•
EP 972828	A1 20000119	EP 1998-401554	19980624
R: AT, BE, CH,	DE, DK, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,
IE, SI, LT,	LV, FI, RO		
CA 2334767	A1 19991229	CA 1999-2334767	19990623
PRIORITY APPLN. INFO.:		EP 1998-401554	A 19980624
		WO 1999-EP4337	W 19990623

The invention relates to a use of a culture medium containing: one or several AB mammalian plasma or sera; a chemical or biol. compound having an antioxidative property and/or differentiating property, such as DMSO, retinoic acid, vitamin, for example vitamin E, or selenium; and/or one or several corticoids for the in vitro hepatitis C virus replication in primary mammalian hepatocytes.

CAPLUS COPYRIGHT 2007 ACS on STN ANSWER 14 OF 27

ACCESSION NUMBER:

2000:755216 CAPLUS

DOCUMENT NUMBER:

133:317537

TITLE:

Hepatitis C virus (HCV) combination therapy,

containing ribavirin in association with antioxidants

Brass, Clifford A.; Glue, Paul W.; Piken, Edward

INVENTOR(S): PATENT ASSIGNEE(S):

Schering Corporation, USA Eur. Pat. Appl., 16 pp.

SOURCE: CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.						KIND DATE					ICAT		· DATE				
	EP	1046	 399											20000418				
		R:						ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
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	WO	2000															00004	
		W:						AU,										
								EE,										
			IS,	JP,	KG,	KR,	KZ,	LC,	LK,	LR,	LT,	LU,	LV,	ΜA,	MD,	MG,	MK,	MN,
								RO,										
								YU,										
		RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZW,	AT,	BE,	CH,	CY,	DE,
			DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,
								GW,										
	BR	2000	00984	40		A		2002	0108		BR 2	2000-	9840			2	0000	118
	HU	2002	0094	2		A2		2002	0729		HU 2	2002-	942		•	2	0000	418
	JP	2002	5422	02		т		2002	1210		JP 2	2000-	6119	35		2	0000	418
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PRIORITY APPLN. INFO.:										2000-					0000			
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Methods are disclosed for treating patients having susceptible viral ΑB infections, especially chronic hepatitis C infection, by administering to the patient a therapeutically effective amount of a combination therapy of interferon- $\alpha$  and ribavirin for a time sufficient to lower HCV-RNA in association with a therapeutically effective amount of an antioxidant for a time sufficient to ameliorate ribavirin-related hemolysis.